

Substitute Form PTO-1449 (Modified)  <b>Information Disclosure Statement by Applicant</b> (Use several sheets if necessary)  (37 CFR §1.98(b))	U.S. Department of Commerce Patent and Trademark Office		Attorney's Docket No. 14174-104US5	Application No. 10/612,179
	Applicant Kreutzer <i>et al.</i>			
	Filing Date July 2, 2003		Group Art Unit 1635	

U.S. Patent Documents							
Examiner Initial	Desig. ID	Document Number	Publication Date	Patentee	Class	Subclass	Filing Date If Appropriate
	A1						

Foreign Patent Documents or Published Foreign Patent Applications								
Examiner Initial	Desig. ID	Document Number	Publication Date	Country or Patent Office	Class	Subclass	Translation	
							Yes	No
	<del>A2</del>	<del>DE 196 18 797</del>	<del>03/23/2000</del>	<del>Germany</del>				<del>X</del>
TV	A3	DE 199 03 713.2	01/30/1999	Germany			X	
TV	A4	DE 199 56 568.6	11/24/1999	Germany			X	
TV	A5	EP 1 144 623 B1	08/28/2002	EPO			X	
	<del>A6</del>	<del>EP 1 214 945</del>	<del>06/19/2002</del>	<del>EPO</del>				<del>X</del>
TV	A7	EP 1 230 375 B1	07/06/2005	EPO				
TV	A8	WO 00/44495	08/03/2000	WIPO				

Other Documents (include Author, Title, Date, and Place of Publication)		
Examiner Initial	Desig. ID	Document
TV	A9	U.S. Provisional Patent Application No. 60/117,335, Li <i>et al.</i> (filed January 28, 1999)
	A10	U.S. Provisional Patent Application No. 60/130,377, Pachuk <i>et al.</i> (filed April 21, 1999)
	A11	Agrawal <i>et al.</i> , "Self-Stabilized Oligonucleotides as Novel Antisense Agents," <u>Delivery Strategies for Antisense Oligonucleotide Therapeutics</u> , Edited by Saghir Akhtar, CRC Press, pp. 105-121 (1995)
	A12	Barber <i>et al.</i> , "Mutants of the RNA-Dependent Protein Kinase (PKR) Lacking Double-Stranded RNA Binding Domain I Can Act as Transdominant Inhibitors and Induce Malignant Transformation," <i>Mol. Cell. Biol.</i> , 15(6):3138-3146 (1995)
	A13	Braich <i>et al.</i> , "Regiospecific Solid-Phase Synthesis of Branched Oligonucleotides. Effect of Vicinal 2',5'- (or 2',3'-) and 3',5'-Phosphodiester Linkages on the Formation of Hairpin DNA," <i>Bioconjug. Chem.</i> , 8:370-377 (1997)
	A14	Brennicke <i>et al.</i> , "RNA editing," <i>FEMS Microbiology Reviews</i> , 23:297-316 (1999)
	A15	Byrom <i>et al.</i> , "Inducing RNAi with siRNA Cocktails Generated by RNase III," <i>TechNotes 10(1)</i> , Ambion, <a href="http://www.ambion.com/techlib/tn/101/4.html">http://www.ambion.com/techlib/tn/101/4.html</a> (2004)
	A16	Chien <i>et al.</i> , "Novel cationic cardiolipin analogue-based liposome for efficient DNA and small interfering RNA delivery <i>in vitro</i> and <i>in vivo</i> ," <i>Cancer Gene Therapy</i> , pp. 1-8 (2004)
	A17	Couzin, "Small RNAs Make Big Splash," <i>Science</i> , 298:2296-2297 (2002)
	A18	Czauderna <i>et al.</i> , "Structural variations and stabilizing modifications of synthetic siRNAs in mammalian cells," <i>Nucleic Acids Res.</i> , 31(11):1-12 (2003)

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TV	A19	Dellweg et al., ed., <u>Römp Lexikon Biotechnologie</u> , p. 354 and p. 673 (1992) (in German)
	A20	Elbashir <i>et al.</i> , "Analysis of gene function in somatic mammalian cells using small interfering RNAs," <i>Methods</i> , 26:199-213 (2002)
	A21	Fallert-Müller, ed., <u>Encyclopedia of Biochemistry</u> , Vol. J-Z, pp. 448-449 (2000) (in German)
	A22	Gryaznov <i>et al.</i> , "Template controlled coupling and recombination of oligonucleotide blocks containing thiophosphoryl groups," <i>Nucleic Acids Res.</i> , 21(6):1403-1408 (1993)
	A23	Hedges, "The Origin and Evolution of Model Organisms," <i>Nature Reviews</i> , 3:838-849 (2002)
	A24	Hornung <i>et al.</i> , "Sequence-specific potent induction of IFN- $\alpha$ by short interfering RNA in plasmacytoid dendritic cells through TLR7," <i>Nature Medicine</i> , 11(3):263-270 (2005)
	A25	Hu-Lieskovan <i>et al.</i> , "Sequence-Specific Knockdown of EWS-FLI1 by Targeted, Nonviral Delivery of Small Interfering RNA Inhibits Tumor Growth in a Murine Model of Metastatic Ewing's Sarcoma" <i>Cancer Res.</i> , 65(19):8984-8992 (2005)
	A26	Hunter <i>et al.</i> , "The characteristics of inhibition of protein synthesis by double stranded ribonucleic acid in reticulocyte lysates," <i>J. Biol. Chem.</i> , 250(2):409-417 (1975)
	A27	"InBase, The Intein Database: The Intein Registry - Inteins Sorted by Species," <a href="http://tools.neb.com/inbase/list.php">http://tools.neb.com/inbase/list.php</a> (database updated on May 22, 2006)
	A28	International Preliminary Examination Report from PCT/DE00/00244
	A29	"Introduction of DNA into Mammalian Cells," <u>Current Protocols in Molecular Biology</u> , Supplement 48, Edited by Frederick M. Ausubel <i>et al.</i> , John Wiley & Sons, Inc., pp. 9.4.7-9.4.8 (1999)
	A30	Judge <i>et al.</i> , "Sequence-dependent stimulation of the mammalian innate immune response by synthetic siRNA," <i>Nat. Biotechnol.</i> , pp. 1-6 (2005) (8 pages of supplementary content included)
	A31	Kennerdell <i>et al.</i> , "Use of dsRNA-Mediated Genetic Interference to Demonstrate that <i>frizzled</i> and <i>frizzled 2</i> Act in the Wingless Pathway," <i>Cell</i> , 95:1017-1026 (1998)
	A32	Kitabwalla <i>et al.</i> , "RNA-Interference - A New Weapon Against HIV and Beyond," <i>N. Engl. J. Med.</i> , 347(17):1364-1367 (2002)
	A33	Lee <i>et al.</i> , "The <i>C. elegans</i> Heterochronic Gene <i>lin-4</i> Encodes Small RNAs with Antisense Complementarity to <i>lin-14</i> ," <i>Cell</i> , 75:843-854 (1993)
	A34	Letter to the International Examining Authority from Gassner & Partner in the prosecution of PCT/DE00/00244 (WO 00/44895), 5 pages (March 28, 2001) (in German)
	A35	Marques <i>et al.</i> , "Activation of the mammalian immune system by siRNAs," <i>Nat. Biotechnol.</i> , 23(11):1399-1405 (2005)
	A36	Martinez <i>et al.</i> , "Single-Stranded Antisense siRNAs Guide Target RNA Cleavage in RNAi," <i>Cell</i> , 110:563-574 (2002)
	A37	McManus <i>et al.</i> , "Gene Silencing in Mammals by Small Interfering RNAs," <i>Nat. Rev. Genet.</i> , 3:737-747 (2002)
	A38	Moss <i>et al.</i> , "The Cold Shock Domain Protein LIN-28 Controls Developmental Timing in <i>C. elegans</i> and Is Regulated by the <i>lin-4</i> RNA," <i>Cell</i> , 88:637-646 (1997)
	A39	Nielsen <i>et al.</i> , "A novel class of conformationally restricted oligonucleotide analogues: synthesis of 2',3'-bridged monomers and RNA-selective hybridization," <i>Chem. Commun.</i> , pp. 825-826 (1997)
✓	A40	PCT/GB00/04404 as filed (November 19, 1999)

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TV	A41	Pegram <i>et al.</i> , "Phase II Study of Receptor-Enhanced Chemosensitivity Using Recombinant Humanized Anti-p185 <sup>HER2/neu</sup> Monoclonal Antibody Plus Cisplatin in Patients With HER2/neu-Overexpressing Metastatic Breast Cancer Refractory to Chemotherapy Treatment," <i>J. Clin. Oncol.</i> , 16(8):2659-2671 (1998)
	A42	Perler, "InBase: the Intein Database," <i>Nucleic Acids Res.</i> , 30(1):383-384 (2002)
	A43	Regalado, "Turning Off Genes Sheds New Light On How They Work," <i>The Wall Street Journal</i> , 4 pages (August 6, 2002)
	A44	Robbins <i>et al.</i> , "Sensing the danger in RNA," <i>Nat. Med.</i> , 11(3):250-251 (2005)
	A45	Schwarz <i>et al.</i> , "Evidence that siRNAs Function as Guides, Not Primers, in the <i>Drosophila</i> and Human RNAi Pathways," <i>Mol. Cell</i> , 10:537-548 (2002)
	A46	Sharp <i>et al.</i> , "RNAi and double-strand RNA," <i>Genes Dev.</i> , 13:139-141 (1999)
	A47	Shi <i>et al.</i> , "A CBP/p300 homolog specifies multiple differentiation pathways in <i>Caenorhabditis elegans</i> ," <i>Genes Dev.</i> , 12:943-955 (1998)
	A48	Sinha, "Large-scale Synthesis. Approaches to Large-scale Synthesis of Oligodeoxynucleotides and their Analogs," <i>Antisense - From Technology to Therapy</i> , Volume 6, Edited by Reimar Schlingensiepen <i>et al.</i> , pp. 29-58 (1997)
	A49	Skipkin <i>et al.</i> , "Psoralen crosslinking between human immunodeficiency virus type 1 RNA and primer (RNA) <sub>3</sub> <sup>Lys</sup> ," <i>Nucleic Acids Res.</i> , 24(3):509-514 (1996)
	A50	Sledz <i>et al.</i> , "Activation of the interferon system by short-interfering RNAs," <i>Nat. Cell Biol.</i> , 5(9):834-839 (2003)
	A51	Soutschek <i>et al.</i> , "Therapeutic silencing of an endogenous gene by systemic administration of modified siRNAs," <i>Nature</i> , 432:173-178 (2004)
V	A52	Strauss, "Candidate 'Gene Silencers' Found," <i>Science</i> , 286:886 (1999)
	A53	Timmons <i>et al.</i> , "Specific interference by ingested dsRNA," <i>Nature</i> , 395:854 (1998)
	A54	Tuschl <i>et al.</i> , "Targeted mRNA degradation by double-stranded RNA in vitro," <i>Genes Dev.</i> , 13:3191-3197 (1999)
	A55	Voinnet <i>et al.</i> , "Systemic signalling in gene silencing," <i>Nature</i> , 389:553 (1997)
	A56	Wargelius <i>et al.</i> , "Double-Stranded RNA Induces Specific Developmental Defects in Zebrafish Embryos," <i>Biochem. Biophys. Res. Commun.</i> , 263:156-161 (1999)
	A57	Waterhouse <i>et al.</i> , "Virus resistance and gene silencing in plants can be induced by simultaneous expression of sense and antisense RNA," <i>Proc. Natl. Acad. Sci. USA</i> , 95:13959-13964 (1998)
	A58	Wess <i>et al.</i> , "Early days for RNAi," <i>BioCentury</i> , 11(12):A1-A8 (2003)
	A59	Wianny <i>et al.</i> , "Specific interference with gene function by double-stranded RNA in early mouse development," <i>Nat. Cell Biol.</i> , 2:70-75 (2000)
	A60	Zamore <i>et al.</i> , "RNAi: Double-Stranded RNA Directs the ATP-Dependent Cleavage of mRNA at 21 to 23 Nucleotide Intervals," <i>Cell</i> , 101:25-33 (2000)
	A61	Zeng <i>et al.</i> , "RNA interference in human cells is restricted to the cytoplasm," <i>RNA</i> , 8:855-860 (2002)

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TV	A62	Zhao <i>et al.</i> , "Double-Stranded RNA Injection Produces Nonspecific Defects in Zebrafish," <i>Dev. Biol.</i> , 229:215-223 (2001)
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